IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

Applicant:

DAGGETT et al.

Serial No.:

08/935,105

Filed:

September 29, 1997

For:

HUMAN N-METHYL-D-ASPARTATE RECEPTOR SUBUNITS, NUCLEIC ACIDS ENCODING SAME AND USES

THEREFOR

Art Unit:

Unassigned

Examiner:

Unassigned

I hereby certify that this paper and the **Statch** papers are being deposited with the United States Postal Service as first class mail in an

envelope addressed to:

Assistant Commissioner for Patents,

Washington, D.C. 20231, on this date.

11/18/97

Date

Nancy V. M¢EĬrath

INFORMATION DISCLOSURE STATEMENT IN ACCORDANCE WITH 37 C.F.R. § 1.97(b) and 1.98

Assistant Commissioner for Patents Washington, D.C. 20231

Dear Sir:

Because this Information Disclosure Statement is filed prior to receipt of a First Office Action on the Merits for the above-captioned application, a fee for filing this statement should not be due. If it is, however, determined that a fee is due, any fees that may be due in connection with filing this paper may be charged to Deposit Account No. 02-4070.

In accordance with the duty of disclosure imposed by 37 C.F.R. §1.56 to inform the Patent Office of all references known by Applicant or Applicant's representative that may be material to the examination of the subject application, Applicant's representative hereby provides this Information Disclosure Statement that is prepared in accordance with 37 C.F.R. §§1.97-1.98. Forms PTO-1449 (10 pages) are provided herewith. Copies of the references listed on the Form PTO-1449 are not provided herewith as they have been previously provided in connection with U.S. Serial No. 08/231,193, which is relied upon for an earlier filing date in accordance with 35 U.S.C. §120.

U.S.S.N. 08/935,105 DAGGETT *et al.* INFORMATION DISCLOSURE STATEMENT

The document CK that is newly cited in connection with this application is attached hereto and is in the English language. Other than as noted below, the documents listed on the Forms PTO-1449 are in the English language. The Japanese patent no. 6014783 (item 0) is in the Japanese language. The foreign patents PCT International Patent Nos. 93/23536, 93/25679, 94/01094, 94/04698, and 95/26401 (items T, V, W, A, and AA respectively) have English language abstracts. Hence, in accordance with the requirements of 37 C.F.R. §1.98, as amended effective March 16, 1992, no further explanation of the listed items are necessary.

The Examiner's attention is directed to reference BI (Hess *et al.*) which is an abstract that published in connection with the 1994 Biophysical Society Annual Meeting. The abstract reports isolation of cDNA clones encoding human N-methyl-D-aspartate receptor subunits, but does not provide the nucleotide sequences of the human clones. A poster presented at the poster session showed a protein sequence comparison among the deduced receptor proteins hNMDAR1A, hNMDAR2A, and hNMDAR2B, but did not provide any nucleotide sequences of the corresponding clones. A copy of the poster is provided in reference BI.

Although these documents are made known to the Patent and Trademark Office in compliance with Applicant's duty of disclosure, such disclosure is not to be construed as an admission by Applicant or Applicant's representative that any of the references, singly or in any combination thereof, is effective as prior art against the subject application. In accordance with 37 C.F.R. §1.97(h), the filing of this Information Disclosure Statement shall not be construed to mean that a search has been made or that no other material information as defined in 37 C.F.R. §1.56(b) exists.

U.S.S.N. 08/935,105 DAGGETT *et al.* INFORMATION DISCLOSURE STATEMENT

Applicant also makes known to the Examiner the following related, copending applications:

U.S.S.N	Inventor	Filing date					
08/052,449 08/480,474 08/486,273 08/940,086 08/940,035	Daggett <i>et al.</i>	4/20/93 6/6/95 6/6/95 9/29/97 9/29/97					
Corresponding Internation	nal Applications						
WO94/24284 4/20/94 G.B. \503689.3 4/20/94 EP 94916547.6-1212 (PUB. NO. 0696320) 4/20/94 AU 68175/94 4/20/94 JP 523578/94 4/20/94 CA 2159106 4/20/94							

Applicant respectfully requests that the Examiner review the foregoing reference and make it of record in the file history of the above-captioned application. ***

Respectfully submitted, BROWN, MARTIN, HALLER & McCLAIN

By:

Stephanie Seidman Registration No. 33,779

Attorney Docket No. 6362-9383D

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FORM PTO-1449 (Modified)

ATTY. DOCKET NO. 6362-9383D

SERIAL NO. 08/935,105

LIST OF PATENTS AND PUBLICATIONS FOR APPLICANT'S INFORMATION DISCLOSURE STATEMENT APPLICANT DAGGETT *et al.*

FILING DATE September 29, 1997

GROUP Unassigned



U.S. PATENT DOCUMENTS

А						ER		DATE	NAME	CLASS	SUB CLASS	FILING DATE
	4	8	3	7	1	4	8	6/6/89	Cregg	435	172.3	10/30/84
В	4	8	5	5	2	3	1	8/8/89	Stroman et al.	435	63	9/25/85
С	4	8	8	2	2	7	9	11/21/89	Cregg	435	68	10/25/85
D	4	9	2	9	5	5	5	5/29/90	Cregg <i>et al.</i>	435	172.3	10/19/87
E	5	0	2	4	9	3	9	6/18/91	Gorman	435	69.1	9/25/87
F	5	0	2	8	7	0	7	7/2/91	Nichols et al.	546	156	11/20/89
G	5	2	0	2	2	5	7	4/13/93	Heinemann et al.	435	252.3	6/21/91
н	5	4	0	1	6	2	9	3/28/95	Harpold <i>et al</i> .	435	6	8/7/90
-	5	4	0	3	4	8	4	4/4/95	Ladner <i>et al.</i>	435	235.1	1/26/93
J	5	4	3	6	1	2	8	7/25/95	Harpold <i>et al</i> .	435	6	1/27/93
	D E F G H -	D 4 E 5 F 5 G 5 H 5	D 4 9 E 5 0 F 5 0 G 5 2 H 5 4 I 5 4	D 4 9 2 E 5 0 2 F 5 0 2 G 5 2 0 H 5 4 0 I 5 4 0	D 4 9 2 9 E 5 0 2 4 F 5 0 2 8 G 5 2 0 2 H 5 4 0 1 I 5 4 0 3	D 4 9 2 9 5 E 5 0 2 4 9 F 5 0 2 8 7 G 5 2 0 2 2 H 5 4 0 1 6 I 5 4 0 3 4	D 4 9 2 9 5 5 E 5 0 2 4 9 3 F 5 0 2 8 7 0 G 5 2 0 2 2 5 H 5 4 0 1 6 2 I 5 4 0 3 4 8	D 4 9 2 9 5 5 5 5 E 5 D 2 4 9 3 9 F 5 D 2 8 7 D 7 G 5 2 D 2 2 5 7 H 5 4 D 1 6 2 9 I 5 4 D 3 4 8 4	D 4 9 2 9 5 5 5 5/29/90 E 5 0 2 4 9 3 9 6/18/91 F 5 0 2 8 7 0 7 7/2/91 G 5 2 0 2 2 5 7 4/13/93 H 5 4 0 1 6 2 9 3/28/95 I 5 4 0 3 4 8 4 4/4/95	D 4 9 2 9 5 5 5 5/29/90 Cregg et al. E 5 0 2 4 9 3 9 6/18/91 Gorman F 5 0 2 8 7 0 7 7/2/91 Nichols et al. G 5 2 0 2 2 5 7 4/13/93 Heinemann et al. H 5 4 0 1 6 2 9 3/28/95 Harpold et al. I 5 4 0 3 4 8 4 4/4/95 Ladner et al.	D 4 9 2 9 5 5 5 5/29/90 Cregg et al. 435 E 5 0 2 4 9 3 9 6/18/91 Gorman 435 F 5 0 2 8 7 0 7 7/2/91 Nichols et al. 546 G 5 2 0 2 2 5 7 4/13/93 Heinemann et al. 435 H 5 4 0 1 6 2 9 3/28/95 Harpold et al. 435 I 5 4 0 3 4 8 4 4/4/95 Ladner et al. 435	D 4 9 2 9 5 5 5 5/29/90 Cregg et al. 435 172.3 E 5 0 2 4 9 3 9 6/18/91 Gorman 435 69.1 F 5 0 2 8 7 0 7 7/2/91 Nichols et al. 546 156 G 5 2 0 2 2 5 7 4/13/93 Heinemann et al. 435 252.3 H 5 4 0 1 6 2 9 3/28/95 Harpold et al. 435 6 I 5 4 0 3 4 8 4 4/4/95 Ladner et al. 435 235.1

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		DOCUMENT NUMBER							COUNTRY	CLASS	SUB CLASS	Tran NO	slation YES
 Κ	0	6	0	0	2	7	8	6/8/94	EP A2				
L	0	6	0	6	7	3	4	7/20/94	EP				
 М	0	6	7	4	0	0	3	9/27/95	EP			÷	
 N	2	2	9	1	6	4	7	1/31/96	GB				
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 Р	9	1	0	6	6	4	8	5/16/91	PCT				
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S	9	3	1	3	4	2	3	7/8/93	PCT				
 Т	9	3	2	3	5	3	6	11/25/93	PCT			•	
 U	9	3	2	4	6	2	9	12/9/93	PCT				

EXAMINER

DATE CONSIDERED

FORM PTO-1449 (Modified)	ATTY. DOCKET NO. SERIAL NO. 6362-9383D 08/935,105				
LIST OF PATENTS AND PUBLICATIONS FOR APPLICANT'S INFORMATION DISCLOSURE	APPLICANT DAGGETT et al.				
STATEMENT	FILING DATE September 29, 1997	GROUP Unassigned			

FOREIGN PATENT DOCUMENTS

		D	осим	ENT 1	NUMBI	ER		DATE	COUNTRY	CLASS	SUB CLASS	Trans NO	slation YES
 V	9	3	2	5	6	7	9	12/23/93	PCT			•	
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 Υ	9	4	0	6	4	2	8	3/31/94	PCT				
Z	9	4	1	1	5	0	1	5/26/94	PCT				
АА	9	5	2	6	4	0	1	10/5/95	PCT			•	

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A	AB	Abbott, NMDA receptor cloned, Trends Pharmacol. Sci. 12:449 (1991)
A	AC	Abbott, NMDA receptor subunit cloned, Trends Pharmacol. Sci. 12:334 (1991)
A	AD	Abe <i>et al.</i> , Molecular characterization of a novel metabotropic glutamate receptor mGluR5 coupled to inositol phosphate/Ca ²⁺ signal transduction, <i>J. Biol. Chem. 267</i> :13361-13368 (1992)
A	ĄΕ	Albin <i>et al.</i> , Abnormalities of striatal projection neurons and <i>N</i> -methyl-D-aspartate receptors in presymptomatic Huntington's Disease, <i>N. Engl. J. Med. 322(18)</i> :1293-1298 (1990)
A	ΑF	Anantharam et al., Combinatorial RNA splicing alters the surface charge on the NMDA receptor, FEBS Lett. 305(1):27-30 (1992)
Δ	٩G	Bahouth et al., Immunological approaches for probing receptor structure and function, Trends Pharmacol. Sci. 12:338-343 (1991)
Д	АН	Barnard, Will the real NMDA receptor please stand up? <i>Trends Pharmacol. Sci. 13</i> :11-12 (1992)
	ΑI	Beal, Mechanisms of excitotoxicity in neurologic diseases, FASEB J. 6:3338-3344 (1992)
Δ	LF	Ben-Ari et al., Protein kinase C modulation of NMDA currents: an important link for LTP induction, <i>Trends Neurosci.</i> 15:333-339 (1992)
А	AK	Black <i>et al.</i> , <i>N</i> -methyl-D-aspartate- or glutamate-mediated toxicity in cultured rat cortical rat cortical neurons is antagonized by FPL 15896AR, <i>J. Neurochem. 65</i> :2170-2177 (1995)

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AL	Bottaro <i>et al,</i> Identification of the hepatocyte growth factor receptor as the c- <i>met</i> proto- oncogene product, <i>Science 251</i> :802-804 (1991)
АМ	Bradford, A rapid and sensitive method for the quantitation of microgram quantities of protein utilizing the principle of protein-dye binding, <i>Anal. Biochem. 72</i> :248 (1976)
AN	Bristow <i>et al.</i> , The glycine/NMDA receptor antagonist R-(+)-HA-966, blocks activation of the mesolimbic dopaminergic system induced by phencyclidine and dizcilpine (MK-801) in rodents, <i>Br. J. Pharmacol.</i> 108:1156-1163 (1993)
AO	Choi, Calcium-mediated neurotoxicity: Relationship to specific channel types and role in ischemic damage, <i>Trends Neurosci.</i> 11(10):465469 (1988)
АР	Choi, Glutamate neurotoxicity and diseases of the nervous system, <i>Neuron</i> 1:623-634 (1988)
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AR	Coyle <i>et al.</i> , Oxidative stress, glutamate, and neurodegenerative disorders, <i>Science</i> 262:689-695 (1993)
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АТ	Dascal, The use of <i>Xenopus</i> oocytes for the study of ion channels, <i>CRC Critical Reviews</i> in <i>Biochemistry 22(4)</i> :317-387 (1987)
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AV	Durand <i>et al.</i> , Cloning of an apparent splice variant of the rat <i>N</i> -methyl-D-aspartate receptor NMDAR1 with altered sensitivity to polyamines and activators of protein kinase C, <i>Proc. Natl. Acad. Sci. USA 89</i> :9359-9363 (1992)
AW	Egebjerg <i>et al.</i> , Intron sequence directs RNA editing of the glutamate receptor subunit GluR2 coding sequence, <i>Proc. Natl. Acad. Sci. USA 91</i> :10270-10274 (1994)
AX	Felder et al., A transfected m1 muscarinic acetylcholine receptor stimulates adenylate cyclase via phosphatidylinisitol hydrolysis, J. Biol. Chem. 264:20356-20362 (1989)
AY	Fisher and Aronson, Characterization of the cDNA and genomic sequence of a G protein γ subunit (γ_5), Mol. Cell. Bio. 12:1585 (1992)
AZ	Foldes <i>et al.</i> , Cloning and sequence analysis of cDNAs encoding human hippocampus <i>N</i> -methyl-D-aspartate receptor subunits: Evidence for alternative splicing, <i>Gene 131</i> :293-298 (1993)

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BB	Gautam <i>et al.</i> , G protein diversity is increased by associations with a variety of <i>y</i> subunits, <i>Proc. Natl. Acad. Sci. USA 87</i> :7973 (1990)
 BC	Gereau and Conn, Multiple presynaptic metabotropic glutamate receptors modulate excitatory and inhibitory synaptic transmission in hippocampal area CA1, <i>J. Neurosci</i> 15(10):6879-6889 (1995)
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 BE	Grimwood <i>et al.</i> , Interactions between the glutamate and glycine recognition sites of the <i>N</i> -methyl-D-aspartate receptor from rat brain, as revealed from radioligand binding studies, <i>J. Neurochem. 60</i> :1729-1738 (1993)
BF	Gubler <i>et al.</i> , A simple and very efficient method for generating cDNA libraries, <i>Gene 25</i> :263-269 (1983)
BG	Gunasekar <i>et al.</i> , NMDA receptor activation produces concurrent generation of nitric oxide and reactive oxygen species: Implication for cell death, <i>J. Neurochem.</i> 65:2016-2021 (1995)
ВН	Gundersen et al., Glutamate and kainate receptors induced by rat brain messenger RNA in Xenopus oocytes, Proc. R. Soc. London Ser. 221:127 (1984)
ВІ	Hess <i>et al.</i> , Cloning, functional expression, and pharmacological characterization of human NMDAR1/NMDAR2 heteromeric receptors, <i>Biophys J., 36(2)</i> :446 (1994) (abstract and poster)
BJ	Hess et al., Biophysical properties of human NMDA receptors stably expressed in mammalian cells, Soc. Neurosci. Abstr. 21:1-3 (1995)
вк	Hoffman, NMDA receptor cloned —— twice! Science 254:801-802 (1991)
BL	Hollman <i>et al.</i> , Zinc potentiates agonist-induced currents at certain splice variants of the NMDA receptor, <i>Neuron 10</i> :943-954 (1993)
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BN	Hurley et al., Isolation and characterization of a cDNA clone for the y subunit of bovine retinal transducin, <i>Proc. Natl. Acad. Sci. USA 81</i> :6948 (1984)
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CF	Linder and Gilman, G proteins, Scientific American 267:56-65 (1992)
CG	Liu <i>et al.</i> , Mutational analysis of the relative orientation of transmembrane helices I and VII in G protein-coupled receptors, <i>J. Biol. Chem.</i> 270(3):19532-19539 (1995)
СН	Lynch <i>et al.</i> , Pharmacological chacterization of heterodimeric NMDA receptors of NR1a and 2B subunits: Differences with receptors formed from NR 1a and 2A, <i>J. Neurochem. 64</i> :1462-1468 (1995)
CI	Masayuki, Human mRNA for key subunit of the N-methyl-D-aspartate receptor, DDBJ database (7/20/93)
CJ	Masu <i>et al.</i> , Sequence and expression of a metabotropic glutamate receptor, <i>Nature</i> 349:760-765 (1991)
СК	Matsui <i>et al.</i> , Functional comparison of D-serine and glycine in rodents: the effect on cloned NMDA receptors and the extracellular concentration, <i>J. Neurochemistry 65:</i> 454-458 (1995)
CL	Mayer, NMDA receptors cloned at last, <i>Nature 354</i> :16-17 (1991)
СМ	Meguro <i>et al.</i> , Functional characterization of a heteromeric NMDA receptor channel expressed from cloned cDNAs, <i>Nature 357</i> :70-74 (1992)
CN	Meldrum, Possible therapeutic applications of antagonists of excitatory amino acid neurotransmitters, <i>Clin. Sci. 68</i> :113-122 (1985)
СО	Meldrum <i>et al.</i> , Excitatory amino acid neurotoxicity and neurodegenerative disease, <i>Trends Pharmacol. Sci. 11</i> :379-387 (1990)
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CO	Monaghan et al., The excitory amino acid receptors: Their classes, pharmacology, and distinct properties in the function of the central nervous system, Ann. Rev. Pharmacol. Toxicol. 29:365-402 (1980)
CR	Monyer et al., Heteromeric NMDA receptors: Molecular and functional distinction of subtypes, Science 256:1217-1221 (1992)
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CV	Nakanishi, Molecular diversity of glutamate receptors and implications for brain function, <i>Science 258</i> :597-602 (1992)
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CY	Ogita et al., A possible role of glutathione as an endogenous agonist at the N-methyl-D-aspartate recognition domain in rat brain, J. Neurochem. 64:1088-1096 (1995)
CZ	Other News to Note, BioWorld Today, 6 (April 15, 1994)
DA	O'Connor et al., Tetanically induced LTP involves a similar increase in the AMPA and NMDA receptor components of the excitory postsynaptic current: Investigations of the involvement of mGlu receptors, <i>J. Neurosci.</i> 15(3):2013-2020 (1995)
	Paoletti and Ascher, Mechanosensitivity of NMDA receptors in cultured mouse central neurons, <i>Neuron</i> 13:645-655 (1995)
DB	Pin et al., Alternative splicing generates metabotropic glutamate receptors inducing different patterns of calcium release in <i>Xenopus</i> oocytes, <i>Neurobiology 89</i> :10331-10335 (1992)
DC	Planells-Cases <i>et al.</i> , Molecular cloning, functional expression, and pharmacological characterization of an <i>N</i> -methyl-D-aspartate receptor subunit from human brain, <i>Proc. Natl. Acad. Sci. USA 90</i> :5057-5061 (1993)
DD	Potter, Sibia to collaborate with Ciba-Geigy, BioWorld Today 3:1 (Sep. 17, 1992)
DE	Reeck et al., "Homology" in proteins and nucleic acids: a terminology muddl; e and a way out of it, Cell 50: 667 (1987)
DF	Rueter et al., Glutamate receptor RNA editing in vitro by enzymatic conversion of adenosine to inosine, Science 267:1491-1494 (1995)
DG	Sakurada <i>et al.</i> , Alteration of Ca ²⁺ permeability and sensitivity to Mg ²⁺ and channel blockers by a single amino acid substitution in the <i>N</i> -methyl-D-aspartate, <i>J. Biol. Chem.</i> 268(1):410-415 (1993)

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